htlp //www.brenda. unickoeln/de./php/resutt

| Αпγ | question? | -> Use | the | BRENDA | Discussion | aroups |
|-----|-----------|--------|-----|--------|------------|--------|
| | | | | | | |

| = | PRINT |
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| | |

| Mark a special word or phrase in this record: | Mark! | |
|---|---------------|----------|
| | All organism | I |
| | Bos taurus | |
| | Gallus gallus | 1 |
| | Homo sapiens | |
| Select one or more organism in this record: | | Submit |

EC NUMBER COMMENTARY

2.4.1.212

RECOMMENDED NAME GeneOntology No.

hyaluronan synthase

GO:0050501

SYSTEMATIC NAME

No entries in this field

| SYNONYMS | ORGANISM | COMMENTARY | LITERATURE |
|----------------------------|----------|------------|------------|
| CHAS2 | | SwissProt | - |
| CHAS3 | _ | SwissProt | * |
| DG42 protein | - | SwissProt | - |
| HA synthase | - | SwissProt | - |
| HuHAS1 | - | SwissProt | - |
| hyaluronan synthethase | - | - | - |
| hyaluronate synthase | - | - | - |
| hyaluronate synthetase | - | - | - |
| hyaluronic acid synthase | - | - | - |
| hyaluronic acid synthetase | - | - | - |
| XHAS1 | - | SwissProt | - |
| XHAS2 | - | SwissProt | - |
| XHAS3 | - | SwissProt | _ |

CAS REGISTRY NUMBER COMMENTARY

39346-43-5

REACTION

COMMENTARY

n UDP-N-acetyl-D-glucosamine + n UDP-D-glucuronate = [beta-N-acetyl-D-glucosaminyl(1->4)beta-D-glucuronosyl (1->3)]n+ 2n UDP

The enzyme from Streptococcus Group A and Group C requires Mg2+. It is highly specific for UDP-GlcNAc and UDP-GlcA; no copolymerization is observed if either is replaced by UDP-Glc, UDP-Gal, UDP-GalNAc or UDP-GalA. Similar enzymes have been found in a variety of organisms

REACTION TYPE ORGANISM COMMENTARY LITERATURE hexosyl group transfer - - -

ORGANISM

COMMENTARY LITERATURE

h

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| Gallus gallus | Swissprot | - | | | |
|---|---------------------|--|---|---------------------------------|-------------------------------|
| Homo sapiens | - | <u>13</u> , <u>15</u> | | | |
| Homo sapiens | Swissprot | - | | | |
| Mus musculus | - | <u>3</u> , <u>6</u> , <u>14</u> | | | |
| Mus musculus | Swissprot | - | | | |
| Paramecium bursaria Chlorella virus | 1 - | <u>5</u> | | | |
| Pasteurella multocida | - | <u>9</u> , <u>11</u> | | | |
| Rattus norvegicus | Swissprot | - | | | |
| Streptococcus equisimilis | - | <u>4</u> , <u>10</u> | | | |
| Streptococcus pyogenes | - | <u>1</u> ,2,8, | <u>10</u> , <u>12</u> | | |
| Streptococcus pyogenes | Swissprot | - | | | |
| Xenopus DG42 | - | 7 | | | |
| Xenopus laevis | Swissprot | - | | | |
| | | | | | |
| SUBSTRATE PRODUCT | REACTION DIAGRAM | ORGANISM | COMMENTARY/ Substrate r:=reversible ir:=irreversible | | COMMENTARY/ LIT Product Pr |
| UDP-N- acetyl-D- glucosamine UDP-D-glucuronate + UDP-D- glucuronate | <u>A</u> | Streptococcus pyogenes | - | 1,2,8, <u>10</u> , <u>12</u> | - |
| UDP-N- acetyl-D- glucosamine UDP-D-glucuronate + UDP-D- glucuronate | <u>A</u> | Mus musculus | - | <u>3</u> , <u>6</u> , <u>14</u> | - |
| UDP-N- acetyl-D- glucosamine UDP-D-glucuronate + UDP-D- glucuronate | <u>A</u> | Streptococcus equisimilis | - | <u>4</u> , <u>10</u> | - |
| UDP-N- acetyl-D- glucosamine UDP-D-glucuronate + UDP-D- glucuronate | <u> </u> | Paramecium bursaria Chlorella virus 1 | - | <u>5</u> | - |
| UDP-N- acetyl-D- glucosamine UDP-D-glucuronate + UDP-D- glucuronate | <u> </u> | Xenopus DG42 | - | 7 | - |
| UDP-N- acetyl-D- glucosamine UDP-D-glucuronate + UDP-D- glucuronate | | Pasteurella multocida | - | 9,11 | - |
| UDP-N- acetyl-D- glucosamine UDP-D-glucuronate + UDP-D- glucuronate | A | Homo sapiens | - | 13 . 15 | - |

Swissprot

h

Bos taurus

b e

e e h f e h

h ec

| NATURAL SUBSTRATE | NATURAL PRODUCT | REACTION DIAGRAM | ORGANISM | COMMENTARY SUBSTRATE | | COMMENTARY PRODUCT | | O (F |
|---|-----------------------|---------------------|--|-------------------------|-----------------------|--------------------|---|---------|
| UDP-N- acetyl-D- glucosamine + UDP-D- glucuronate | UDP-D- glucuronate | A | Streptococcus pyogenes | - | 1,2,8,10, 12 | - | - | |
| UDP-N- acetyl-D- glucosamine + UDP-D- glucuronate | UDP-D- glucuronate | <u> </u> | Mus musculus | - | 3,6, <u>14</u> | - | - | |
| UDP-N- acetyl-D- glucosamine + UDP-D- glucuronate | UDP-D- glucuronate | <u>&</u> | Streptococcus equisimilis | - | <u>4</u> , <u>10</u> | - | - | |
| UDP-N- acetyl-D- glucosamine + UDP-D- glucuronate | UDP-D- glucuronate | <u> </u> | Paramecium bursaria Chlorella virus 1 | - | <u>5</u> | - | - | |
| UDP-N- acetyl-D- glucosamine + UDP-D- glucuronate | UDP-D- glucuronate | <u> </u> | Xenopus DG42 | - | 7 | - | - | |
| UDP-N- acetyl-D- glucosamine + UDP-D- glucuronate | UDP-D- glucuronate | <u>A</u> | Pasteurella multocida | - | <u>9</u> , <u>11</u> | - | - | |
| UDP-N- acetyl-D- glucosamine + UDP-D- glucuronate | UDP-D- glucuronate | <u>A</u> | Homo sapiens | - | <u>13</u> , <u>15</u> | - | - | |

COFACTOR ORGANISM COMMENTARY LITERATURE IMAGE

No entries in this field

| METAL IONS | ORGANISM | COMMENTARY | LITERATURE |
|---------------|-------------------------------------|--|------------|
| Co2+ | Paramecium bursaria Chlorella viru | s 2% as effective as Mn2+ at similar concentrations <5> | <u>5</u> |
| Mg2+ | Paramecium bursaria Chlorella viru. | s 20% as effective as Mn2+ at similar concentrations <5> | <u>5</u> |
| M g2+ | Xenopus DG42 | - | <u>7</u> |
| Mn2+ | Paramecium bursaria Chlorella viru | s essential for activity <5> | <u>5</u> |
| Mn2+ | Pasteurella multocida | - | <u>9</u> |
| INHIBITORS | ORGANISM COM | MMENTARY LITERATURE IMAGE | |
| N-ethylmalei | mide Streptococcus pyogenes | - <u>12</u> <u>● 2D-image</u> | |

ACTIVATING COMPOUND ORGANISM COMMENTARY LITERATURE IMAGE

h be e h f e h ec

No entries in this field

| KM VALUE [mM] | KM VALUE [mM] Maximum | SUBSTRATE | ORGANISM | COMMENTARY | LITERATURE | IMAGE |
|---------------------------|-----------------------------|--------------------------------|---------------------------|---|------------|------------------------|
| additional information | - | more | Mus musculus | values for other substrate concentrations <6> | <u>6</u> | - |
| 0.03 | - | UDP-D- glucuronate | Mus musculus | HAS2 <6> | <u>6</u> | <u>2D-</u> image |
| 0.03 | - | UDP-D- glucuronate | Mus musculus | HAS3 <6> | <u>6</u> | ● <u>2D-</u> image |
| 0.04 | - | UDP-D- glucuronate | Streptococcus pyogenes | - | <u>10</u> | ● <u>2D-</u> image |
| 0.05 | - | UDP-D- glucuronate | Streptococcus equisimilis | - | <u>10</u> | ● <u>2D-</u> image |
| 0.06 | - | UDP-D- glucuronate | Xenopus DG42 | - | <u>7</u> | ● <u>2D-</u> image |
| 0.07 | - | UDP-D- glucuronate | Mus musculus | HAS1 <6> | <u>6</u> | ● <u>2D-</u> image |
| 0.14 | - | UDP-D- glucuronate | Pasteurella multocida | - | <u>11</u> | ● <u>2D-</u> image |
| 0.06 | - | UDP-N-acetyl-D- glucosamine | Streptococcus equisimilis | - | <u>10</u> | ● <u>2</u> D- image |
| 0.08 | - | UDP-N-acetyl-D- glucosamine | Mus musculus | HAS3 <6> | <u>6</u> | ● <u>2D-</u> image |
| 0.11 | - | UDP-N-acetyl-D- glucosamine | Mus musculus | HAS2 <6> | <u>6</u> | ● <u>2D-</u> image |
| 0.14 | - | UDP-N-acetyl-D- glucosamine | Streptococcus pyogenes | - | 10 | ● <u>2D-</u> image |
| 0.16 | - | UDP-N-acetyl-D- glucosamine | Pasteurella multocida | - | 11 | ● 2D- image |
| 0.23 | - | UDP-N-acetyl-D- glucosamine | Xenopus DG42 | - | Z | <u>2D-</u> image |
| 0.79 | - | UDP-N-acetyl-D- glucosamine | Mus musculus | HAS1 <6> | <u>6</u> | ● <u>2D-</u> image |
| | | | | | | |

Ki VALUE [mM] Ki VALUE [mM] Maximum INHIBITOR ORGANISM COMMENTARY LITERATURE IMAGE No entries in this field

TURNOVER NUMBER

TURNOVER NUMBER MAXIMUM

SUBSTRATE ORGANISM COMMENTARY LITERATURE IMAGE

No entries in this field

[µM/min/mg]

SPECIFIC ACTIVITY SPECIFIC ACTIVITY MAXIMUM ORGANISM COMMENTARY LITERATURE

No entries in this field

| pH OPTIMUM | pH MAXIMUM | ORGANISM | COMMENTARY | LITERATURE |
|------------|------------|---------------------------------------|------------|------------|
| 7.6 | 8.1 | Xenopus DG42 | - | 7 |
| 7.2 | - | Paramecium bursaria Chlorella virus 1 | _ | <u>5</u> |

h e h f b e h ec

pH RANGE pH RANGE MAXIMUM ORGANISM COMMENTARY LITERATURE

7 8.4 Xenopus DG42 - 7

TEMPERATURE OPTIMUM TEMPERATURE OPTIMUM MAXIMUM ORGANISM COMMENTARY LITERATURE
No entries in this field

TEMPERATURE RANGE TEMPERATURE MAXIMUM ORGANISM COMMENTARY LITERATURE

No entries in this field

SOURCE TISSUE ORGANISM COMMENTARY LITERATURE

breast adenocarcinoma cell Mus musculus B6-cell line <14> 14

glioma cell Homo sapiens cell line <13> 13

keratinocyte Homo sapiens - <u>15</u>

 LOCALIZATION
 ORGANISM
 COMMENTARY
 GeneOntology No.
 LITERATURE

 membrane
 Streptococcus pyogenes
 enzyme is predicted to be an integral membrane protein <1>
 GO:0016020
 1,2,8,12

 membrane
 Mus musculus
 GO:0016020
 3,6

MOLECULAR WEIGHT ACCESSION NO. OF **ENTRY NAME ORGANISM SOURCE** Sequence CODE AA [Da] Show 551 63685 O57427 pBLAST HAS2 XENLA Xenopus laevis Swissprot Sequence Show Q92819 pBLAST HAS2_HUMAN Homo sapiens 552 63566 Swissprot Sequence Show O97711 pBLAST HAS2_BOVIN Bos taurus 552 63459 Swissprot Sequence Show P70312 pBLAST HAS2_MOUSE Mus musculus 552 63492 Swissprot Sequence **₾** Show O35776 pBLAST HAS2_RAT Rattus norvegicus 552 63534 **Swissprot** Sequence **♦** Show P13563 pBLAST HAS1 XENLA Xenopus laevis 588 68522 Swissprot Sequence Streptococcus **७** Show Q54865 pBLAST HASA_STRPY 419 47886 Swissprot pyogenes Sequence Show O00219 pBLAST HAS3_HUMAN Homo sapiens 553 63070 Swissprot Sequence Show O08650 pBLAST HAS3_MOUSE Mus musculus 554 63338 Swissprot Sequence **७** Show Q61647 pBLAST HAS1_MOUSE Mus musculus 583 65544 **Swissprot** Sequence Show Q92839 pBLAST HAS1_HUMAN Homo sapiens 578 64884 **Swissprot** Sequence Show O57424 pBLAST HAS2_CHICK Gallus gallus 552 63744 Swissprot Sequence

PDB ORGANISM

No entries in this field

| MOLECULAR WEIGHT | MOLECULR WEIGHT MAXIMUM | ORGANISM | COMMENTARY | LITERATURE |
|---------------------|-------------------------|---------------------------|-------------------------------|------------|
| 66000 | • | Homo sapiens | gel filtration <13> | <u>13</u> |
| 52000 | <u>u</u> | Mus musculus | SDS-PAGE <14> | <u>14</u> |
| 48000 | • | Mus musculus | northern blot <3> | <u>3</u> |
| 47780 | · - | Streptococcus pyogenes | calculation from sequence <8> | <u>8</u> |
| 47780 | - | Streptococcus equisimilis | calculation from sequence <4> | <u>4</u> |
| 42000 | - | Streptococcus pyogenes | SDS-PAGE <1,2> | 1.2 |
| 42000 | - | Streptococcus equisimilis | - | <u>4</u> |

SUBUNITS ORGANISM COMMENTARY LITERATURE

No entries in this field

POSTTRANSLATIONAL MODIFICATION ORGANISM COMMENTARY LITERATURE

No entries in this field

Crystallization/COMMENTARY ORGANISM LITERATURE

No entries in this field

PH STABILITY PH STABILITY MAXIMUM ORGANISM COMMENTARY LITERATURE

No entries in this field

TEMPERATURE STABILITY TEMPERATURE STABILITY MAXIMUM ORGANISM COMMENTARY LITERATURE

No entries in this field

GENERAL STABILITY ORGANISM LITERATURE

No entries in this field

ORGANIC SOLVENT ORGANISM COMMENTARY LITERATURE

No entries in this field

OXIDATION STABILITY ORGANISM LITERATURE

No entries in this field

STORAGE STABILITY ORGANISM COMMENTARY LITERATURE

4°C, Na-phosphate buffer,10% glycerol, 96 h, 18% Homo sapiens - 13

Purification/COMMENTARY ORGANISM LITERATURE

partial <13> Homo sapiens 13

Cloned/COMMENTARY ORGANISM LITERATURE
expression in COS-1 cells and rat 3Y1 fibroblasts <6> Mus musculus

6

expression in Escherichia coli <1,2,8,10,12> Streptococcus pyogenes 1, 2, 8, 10, 12

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| expression in Escherichia coli <4,10> | Streptococcus equisimilis | <u>4</u> , <u>10</u> |
|---------------------------------------|---------------------------------------|----------------------|
| expression in Escherichia coli <5> | Paramecium bursaria Chlorella virus 1 | <u>5</u> |
| expression in Escherichia coli <9,11> | Pasteurella multocida | <u>9</u> , <u>11</u> |
| expression in yeast <7> | Xenopus DG42 | <u>7</u> |

| ENGINEERING | ORGANISM | COMMENTARY | LITERATURE |
|-------------|------------------------|--|------------|
| D196N | Pasteurella multocida | mutants possess UDP-D-glucuronate-transferase activity <11> | <u>11</u> |
| D477K | Pasteurella multocida | mutants possess UDP-N-acetyl-D-glucosamine-transferase activity <11> | 11 |
| more | Streptococcus pyogenes | variety of cystein mutatants <12> | <u>12</u> |

Renatured/COMMENTARY ORGANISM LITERATURE

No entries in this field

APPLICATION ORGANISM COMMENTARY LITERATURE

No entries in this field

DISEASE TITLE OF PUBLICATION LINK TO PUBMED

No entries in this field

| REF. | AUTHORS | TITLE | JOURNAL | VOL. | PAGES | YEAR | ORGANISM | COMMENTARY | LINK TO PUBME! |
|----------|---|--|-------------------|------|-----------------|------|---------------------------|------------|----------------|
| 1 | DeAngelis, P.L.; Papaconstantinou, J.; Weigel, P.H. | Molecular cloning, identification and sequence of the hyaluronan synthase gene from Group A Streptococcus pyogenes | J. Biol. Chem. | 268 | 19181- 19184 | 1993 | Streptococcus pyogenes | - | ● <u>PubM</u> |
| <u>2</u> | DeAngelis, P.L.; Weigel, P.H. | Immunochemical confirmation of the primary structure of streptococcal hyaluronan synthase and synthesis of high molecular weight product by the recombinant enzyme | Biochemistry | 33 | 9033- 9039 | 1994 | Streptococcus pyogenes | - | ● <u>PubM</u> |
| <u>3</u> | Spicer, A.P.; Augustine, M.L.; McDonald, J.A. | Molecular cloning and characterization of a putative mouse hyaluronan synthase Molecular | J. Biol. Chem. | 271 | 23400- 23406 | 1996 | Mus musculus | - | - |
| | | cloning, expression, and characterization | | | | | | | |

| <u>4</u> | Kumari, K.; Weigel, P.H. | of the authentic hyaluronan synthase from group C Streptococcus equisimilis | J. Biol. Chem. | 272 | 32539- 32546 | 1997 | Streptococcus equisimilis | - | <u>Pub</u> M |
|-----------|--|---|-------------------|-----|-----------------|------|--|---|---------------|
| <u>5</u> | DeAngelis, P.L.; Jing, W.; Graves, M.V.; Burbank, D.E.; Van Etten, J.L. | Hyaluronan synthase of chlorella virus PBCV-1 | Science | 278 | 1800- 1803. | 1997 | Paramecium bursaria Chlorella virus 1 | | ● <u>PubM</u> |
| <u>6</u> | Itano, N.; Sawai, T.; Yoshida, M.; Lenas, P.; Yamada, Y.; Imagawa, M.; Shinomura, T.; Hamaguchi, M.; Yoshida, Y.; Ohnuki, Y.; Miyauchi, S.; Spicer, A.P.; McDonald, J.A.; Kimata, K. | Three isoforms of mammalian hyaluronan synthases have distinct enzymatic properties | J. Biol. Chem. | 274 | 25085- 25092 | 1999 | Mus musculus | - | - |
| Z | Pummill, P.E.; Achyuthan, A.M.; DeAngelis, P.L. | Enzymological characterization of recombinant Xenopus DG42, a vertebrate hyaluronan synthase | J. Biol. Chem. | 273 | 4976- 4981 | 1998 | Xenopus DG42 | - | - |
| <u>8</u> | Tlapak-Simmons, V.L.; Baggenstoss, B.A.; Clyne, T.; Weigel, P.H. | Purification and lipid dependence of the recombinant hyaluronan synthases from Streptococcus pyogenes and Streptococcus equisimilis | J. Biol. Chem. | 274 | 4239- 4245 | 1999 | Streptococcus pyogenes | - | - |
| <u>9</u> | DeAngelis, P.L. | Molecular directionality of polysaccharide polymerization by the Pasteurella multocida hyaluronan synthase | J. Biol. Chem. | 274 | 26557- 26562 | 1999 | Pasteurella multocida | - | ● <u>PubM</u> |
| <u>10</u> | Tlapak-Simmons, V.L.; Baggenstoss, B.A.; Clyne, T.; Weigel, P.H. | Purification and lipid dependence of the recombinant hyaluronan synthases from Streptococcus pyogenes and Streptococcus equisimilis | J. Biol. Chem. | 274 | 4239- 4245 | 1999 | Streptococcus pyogenes, Streptococcus equisimilis | _ | - |
| | | Dissection of the two transferase activities of the | | | | | | | |

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| <u>11</u> | Jing, W.; DeAngelis, P.L. | Pasteurella multocida hyaluronan synthase: two active sites exist in one polypeptide | Glycobiology | 10 | 883- 889 | 2000 | Pasteurella multocida | - | - |
|-----------|--|--|------------------------------|------|---------------|------|---------------------------|---|---------------|
| <u>12</u> | Heldermon, C.D.; Tlapak-Simmons, V.L.; Baggenstoss, B.A.; Weigel, P.H. | Site-directed mutation of conserved cysteine residues does not inactivate the Streptococcus pyogenes hyaluronan synthase | Glycobiology | 11 | 1017- 1024 | 2001 | Streptococcus pyogenes | - | <u>● PubM</u> |
| <u>13</u> | Asplund, T.; Brinck, J.; Suzuki, M.; Briskin, M.J.; Heldin, P. | Characterization of hyaluronan synthase from a human glioma cell line | Biochim. Biophys. Acta | 1380 | 377- 388 | 1998 | Homo sapiens | - | - |
| <u>14</u> | Klewes, L.; Prehm, P. | Intracellular signal transduction for serum activation of the hyaluronan synthase in eukaryotic cell lines | J. Cell.Physiol. | 160 | 539- 544 | 1994 | Mus musculus | - | ●_PubM |
| <u>15</u> | Sayo, T.; Sugiyama, Y.; Takahashi, Y.; Ozawa, N.; Sakai, S.; Ishikawa, O.; Tamura, M.; Inoue, S. | Hyaluronan synthase 3 regulates hyaluronan synthesis in cultured human keratinocytes | J. Invest. Dermatol. | 118 | 43-48 | 2002 | Homo sapiens | - | ● <u>PubM</u> |

LINKS TO OTHER DATABASES (specific for EC-Number 2.4.1.212)

ExPASy

Online Mendelian Inheritance in Man

KEGG

NCBI: PubMed, Protein, Nucleotide, Structure, Genome, OMIM, Domains

IUBMB Enzyme Nomenclature

WIT database

EMP Project

PDB database(3D structure)

PROSITE Database of protein families and domains

SYSTERS

Protein Mutant Database

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b e

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